# **CPQCC** Hypothermia Webinar

December 1, 2021 12pm – 1pm



### **CPQCC Hypothermia Webinar** AGENDA

TIME	TOPIC	SPEAKER
12pm – 12:02pm	Welcome and Intros	Henry Lee, MD
12:02- 12:17pm	CPQCC Early Screening and Identification of Candidates for Neonatal Therapeutic Hypothermia Toolkit	CPQCC Hypothermia Toolkit Author – Dongli Song, MD, PhD
12:17pm – 12:32pm	Current updates on HIE management	CPQCC Hypothermia Toolkit Author – Krisa Van Meurs, MD
12:32pm – 12:47pm	Demo of the CoolTool & NeoCool	<u>CoolTool</u> – Tom Shimotake, MD <u>NeoCool</u> – Mary Harbert, MD
12:47pm – 1:00pm	Q&A	Moderator(s) Henry Lee & Priya Jeegatheesan, MD



### Webinar Logistics

- Attendees are automatically muted upon entry
- The "chat" function has been disabled. Please utilize the Q&A box if you are having technical difficulties and to submit any questions you have for the presenters. We will answer as many questions as possible during the Q&A portion of the webinar.
- The slides and webinar recording will be sent out after the webinar and will also be posted on the CPQCC website.



CPQCC Early Screening and Identification of Candidates for Neonatal Therapeutic Hypothermia Toolkit

Dongli Song, MD, PhD Stay Informed Annual Improvement Palooza Annual Data Training Annual Reports FAQs Glossary Connect With Us

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### Early Screening and Identification of Candidates for Neonatal Therapeutic Hypothermia Toolkit

Therapeutic hypothermia, when initiated within six hours of birth, has been shown to significantly improve survival and neurodevelopmental outcomes in neonates with moderate to serve hypoxic-ischemic encephalopathy (HIE). However, not every baby who might benefit from cooling therapy is identified or referred to a regional cooling center in a timely fashion. Early identification of the risk factors for perinatally-acquired asphyxia and recognition of the signs and symptoms of neonatal encephalopathy are challenging even for the most experienced neonatologists.

- To recognize that these are **screening criteria only**, meant to improve early identification of at-risk babies who might warrant closer assessment.
- They are intentionally designed with more inclusive criteria and are **NOT by themselves qualifying criteria for cooling therapy.**

DOWNLOAD TOOLKIT »

#### Priya Jegatheesan Anna Morgan Thomas Shimotake Dongli Song Krisa Van Meurs

Author:

Resource Category: QI Toolkit

Date: February 2015

Related Links: HIE Calculator (Cool Tool)



### Objective

To provide a **strategic approach** and **reliable tools** to assist birth hospitals in the **timely identification** of newborns that are potential candidates for therapeutic hypothermia.



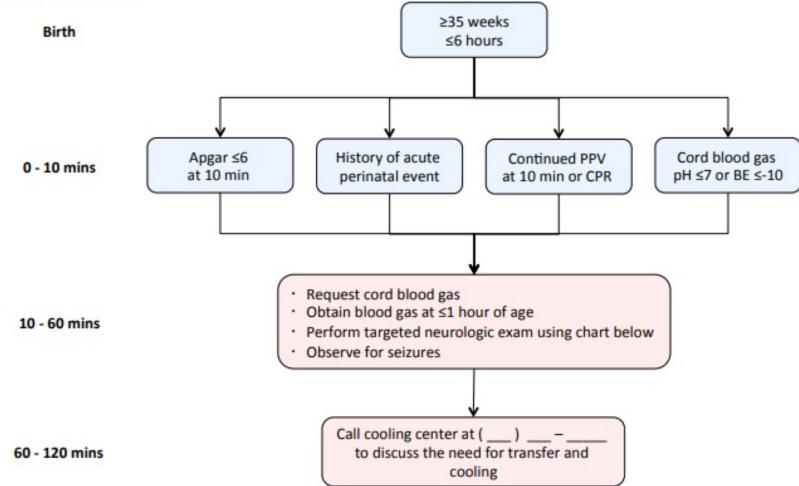
### **Screening Criteria**

 $\geq$  35 weeks gestational age and  $\leq$  6 hours old AND any one of the following must also be present:

- 1) History of acute perinatal event and risk factors:
  - placental abruption, cord prolapse, uterine rupture and fetal bradycardia prolonged rupture of membranes, abnormal fetal heart rate tracings, thick meconium stained fluid, tight nuchal cord, and failed vacuum delivery.
- 1) Apgar  $\leq 6$  at 10 minutes
- 2) Continued need for PPV for 10 minutes or history of CPR
- 3) Venous or arterial cord gas or baby blood gas (within 1 hour of life) with pH  $\leq$  7 or BE  $\leq$  -10

### Appendix A

<u>Screening Criteria for Evaluation of Risk for Neonatal Encephalopathy (NE)</u> Goal timeline





# Neurological Exam

#### Level of encephalopathy

	Mild	Moderate	Severe	
Level of consciousness	Irritable / hyperalert	Lethargic / obtunded	Stupor / coma	
Spontaneous activity	Normal / increased	Decreased	No activity	
Posture	Normal	Distal flexion / Complete extension	Decerebrate	
Tone	Normal / increased	Hypotonic	Flaccid	
Primitive reflexes				
Suck	Normal	Weak	Absent	
Moro	Normal	Incomplete	Absent	
Deep tendon reflexes	Mildly brisk	Brisk	Suppressed	
Autonomic system			Deviated dilated app	
Pupils	Normal	Constricted	Deviated, dilated, non-	
Heart rate	Increased	Bradycardic	reactive Variable	
Seizures	None	Common	Common	



### **Passive Cooling**

- Document Regional Cooling Center contacted and decision made to initiate passive cooling for those determined to be a candidate for cooling.
- 2. Turn radiant warmer off and leave infant uncovered, except diapers.
- 3. Monitor core/rectal temperature continuously (if equipped) or every 15 minutes using a lubricated digital thermometer carefully inserted 2 cm into rectum. If core temperature monitoring cannot be done safely or is not available, monitor axillary temperatures every 15 minutes. Record temperatures on flow sheet (see Appendix H).
- 4. Allow temperature to fall to target temperature ranges:
  - 1. Target rectal temperature is 33-34°C or 91.4-93.2°F.
  - 2. Target axillary temperature is 32-33°C or 89.6-91.4F.

### Passive Cooling -cont.

- Avoid overcooling. When the rectal temp reaches 33 °C (91.4 °F) or axillary temp 32 °C (89.6 °F), turn warmer back on to lowest setting or covering patient with clear plastic (avoid face).
- If rectal temp continues to fall quickly or remains < 33 °C (91.4 °F) or axillary temp < 32 °C (89.6 °F), increase warmer setting. Recheck temperature until recovered.</li>
- Avoid overheating. Minimize big changes in heater settings that may result in overcorrections.
- 8. Monitor vital signs, electrolytes and glucose levels closely.
- 9. If administering respiratory support, avoid hyperoxia and iatrogenic hyperventilation.
- 10. Keep patient comfortable and adequately sedated (i.e., avoid shivering).



### Neonates Who Do Not Qualify for Cooling

#### 1. Maintain communication with regional cooling center

a. Discuss management and plan if significant clinical changes develop.

#### 2. If heat sources were removed/cooling was initiated, slowly begin rewarming

- a. Document time of lowest temperature and source (e.g., axillary vs. rectal).
- b. Rewarm with target rate of approximately 0.5 °C /hour. Avoid overheating.

#### 3. Monitor temperature periodically

- a. Target rectal/core temp = **36.5°C** (97.7°F) or axillary/skin temp = **36.0°C** (96.8°F).
- b. Check temperature periodically (e.g., hourly for first 6 hours).

#### 4. Check glucose and electrolyte levels.

- a. Fluctuations may be seen check Glucose levels. Avoid hypoglycemia
- **b.** Consider maintaining higher normal target glucose levels (e.g., >50mg/dl)
- c. Consider checking Ca, K, Mg levels. Maintain within normal ranges.

#### 5. Obtain follow-up blood gases to confirm acidosis resolving

a. If acidosis persists, work-up other causes or discuss with neonatologist.

### Neonates Who Do Not Qualify for Cooling – Cont.

#### **6.** Repeat neurologic examination (see appendix B)

- a. Document initial neurologic exam.
- b. Repeat neurologic exam (e.g., after 1-3 hours) if clinically indicated.
- c. Document neurologic exam at time of discharge.

#### 7. If initial acidosis severe, consider delaying enteral feeds (NPO) until improved

- a. Depends upon severity of clinical presentation. Discuss with neonatologist.
- b. May require initiation of maintenance IVF fluids.

#### 8. Avoid iatrogenic hyperventilation and hyperoxygenation

- a. Normal pCO2 levels (35-45 mmHg) compensatory hyperventilation may be seen.
- b. Normal PaO2 levels (60-100mmHg) and oxygen saturations (<94-98%).
- 9. Consider ordering baseline labs:
  - a. CBC, platelets and Blood cultures.
  - b. Start antibiotics if appropriate.

# Screening for HIE & Active Cooling - QI

Year	Live birth (inborn) n	Screened n % live birth	NICU n % live birth (% evaluated)	Active Cooled n % live birth (% evaluated in NICU)
2008-2018	42611	1824 4.3%	326 0.78% (18%)	66 <b>0.15%</b> (20%)



Updated 2/12/19

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# Screening for HIE & Active Cooling - QI

Year	Live birth (inborn) n	Screened n % live birth	NICU n % live birth (% evaluated)	Active Cooled n % live birth (% evaluated in NICU)
2008-2018 SCVMC	42611	1824 4.3%	326 0.78% (17.8%)	66 0.15% (1.5 per 1000)
2008-2015 NC Kaiser	44 572	197 (4.4 per 1000) BE -<12		45 0.1% (1 per 1000)

SANTA CLARA VALLEY MEDICAL CENTER Hospital & Clinics

Updated 2/12/19

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Current updates on HIE management

Krisa Van Meurs, MD

# Current Updates on HIE Management Outline

Randomized Clinical Trial results (completed)

- NICHD Late Hypothermia RCT
- NeoLEV RCT

Randomized Clinical Trials (updates)

- NICHD Preemie Hypothermia RCT
- HEAL RCT
- TIME RCT

Recently presented or published studies

- Seizures during rewarming
- AEDs after discharge
- Trophic feeds during cooling

Resources

### NICHD RCT "Late" hypothermia for HIE

Study design: Multi-center, randomized, non-masked, Bayesian analysis

Eligibility: 6-24 hours of age with evidence of moderate or severe encephalopathy

Intervention: Whole body cooling to esophageal temperature 33.5° C x 96 hours or control

1° outcome: Death or moderate/severe disability at 18-22 months of age

Sample size: 168

Eunice Kennedy Shriver

Laptook A et al. JAMA 2017



### Should a baby be cooled after 6 hours?

How this happens:

- Arrival at a cooling center after 6 hrs of age
- Progress from stage I to II/III encephalopathy after 6 hrs of age
- Are not recognized to qualify until after 6hrs of age
- Cooling cannot be initiated within 6 hours of age (equipment or personnel not available)

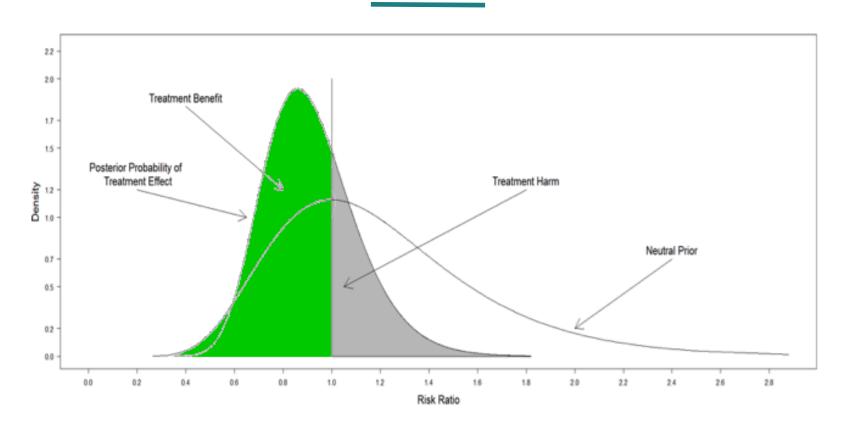
All Cooling Centers should work with their regional hospitals to screen and identify infants for cooling at <6 hours of age so take advantage of the benefit of cooling on death and disability.

### Adjusted Risk Ratios (aRR)\* for primary and secondary outcomes: Bayesian

		Cooled (n=78)		cooled =79)	aRR (Neutral prior)	Probability Benefit (Neutral prior)	
Primary	n	%	n	%	95% credible intervals	aRR<1.0	aRR<0.9
Death or mod/sev disability	19	24.4	22	27.9	.86 (.59-1.29)	.77	.58
Secondary							
Death	9	11.5	9	11.4	.86 (0.53-1.44)	.73	.57
Severe disability	9	11.5	12	15.2	.88 (0.51-1.49)	.68	. 53
Moderate disability	1	1.3	1	1.3			
Mild disability	16	20.5	12	15.2	1.18 (0.72-1.92)	.25	.13



### **Probability of treatment benefit**



**Conclusion:** 77% of newborns cooled between 6-24 hours will have some benefit.

Laptook A et al. JAMA 2017

### NEOLEV2

### Levetiracetam Versus Phenobarbital for Neonatal Seizures: A Randomized Controlled Trial

Cynthia Sharpe, MBChB,<sup>a,b</sup> Gail E. Reiner, DNP,<sup>b</sup> Suzanne L. Davis, MBChB, PhD,<sup>a</sup> Mark Nespeca, MD,<sup>b</sup> Jeffrey J. Gold, MD, PhD,<sup>b</sup> Maynard Rasmussen, MD,<sup>c</sup> Rachel Kuperman, MD,<sup>d</sup> Mary Jo Harbert, MD,<sup>e</sup> David Michelson, MD,<sup>f</sup> Priscilla Joe, MD,<sup>g</sup> Sonya Wang, MD,<sup>b</sup> Neggy Rismanchi, MD, PhD,<sup>b</sup> Ngoc Minh Le, MD,<sup>k</sup> Andrew Mower, MD,<sup>h</sup> Jae Kim, MD,<sup>1</sup> Malcolm R. Battin, MBChB,<sup>m</sup> Brian Lane, MD,<sup>i</sup> Jose Honold, MD,<sup>i</sup> Ellen Knodel, RCP,<sup>i</sup> Kathy Arnell, RN,<sup>k</sup> Renee Bridge, BSN, RN,<sup>1</sup> Lilly Lee, BA,<sup>j</sup> Karin Ernstrom, MS,<sup>n</sup> Rema Raman, PhD,<sup>n</sup> Richard H. Haas, MB, BChir,<sup>b</sup> FOR THE NEOLEV2 INVESTIGATORS

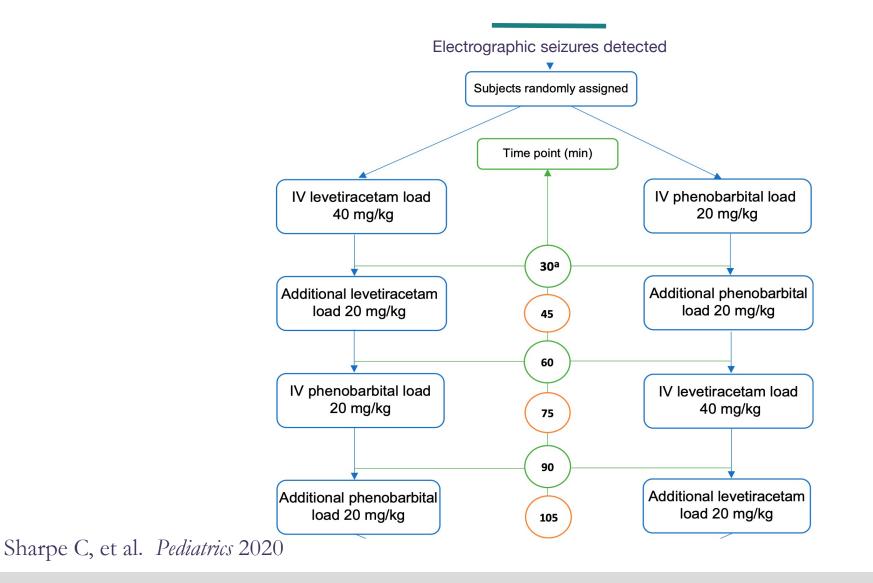
The objective of this study was to determine the efficacy and safety of levetiracetam compared with phenobarbital as a firstline treatment of neonatal seizures.

The primary outcome was complete seizure freedom for 24 hours.

PEDIATRICS Volume 145, number 6, June 2020:e20193182



### **NEOLEV2 - Design**



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### **NEOLEV2 - Results**

	Phenobarbital n=30	Levetiracetam n = 53	P value	RR (95% CI)
Primary outcome measure 24h seizure cessation rate	80%	28%	< 0.001	0.35 (0.22-0.56)
Secondary outcome measures 48h seizure cessation rate 1h seizure cessation rate	64% 93%	17% 49%	<0.001 <0.001	0.26 (0.13-0.53) 0.53 (0.39-0.7)
Subanalysis of patients with HIE treated with hypothermia 24h seizure cessation rate	90%	35%	0.014	0.39 (0.2-0.77)

Sharpe C, et al. *Pediatrics* (2020)



### **NEOLEV 2 - Conclusions**

- Phenobarbital 20-40 mg/kg has a greater efficacy than levetiracetam 40-60 mg/kg.
- More adverse events occurred with phenobarbital, but the difference was not statistically significant.
- Higher dose studies of levetiracetam are warranted given the increased efficacy with dose escalation and the excellent safety profile
- Definitive studies with long-term outcome measures are needed.

Sharpe C, et al. *Pediatrics* (2020)



# Therapeutic hypothermia for HIE in premature infants 33-35 weeks gestation

- Purpose:To determine if cooling benefits infants 33-35 weeks gestation with<br/>moderate to severe HIE
- Methods:Randomized controlled trial whole body hypothermia with esophageal<br/>temperature 33.5° C for 72 hours
- **1° outcome:** Death or moderate to severe neurodevelopmental impairment at 18-22 months
- Sample size: 168, enrollment completed, in follow-up phase
- PIs:Roger Faix, MD and Abbot Laptook, MDNICHD Neonatal Research Network



### Need for cooling plus therapies

	Controls	Cooled
Death or moderate to severe disability	62-83%	44-55%
Death	27-57%	24-38%
Cerebral palsy	30-48%	19-33%

**Conclusion:** Additional neuroprotective strategies are needed to further reduce mortality and morbidity.

Shankaran S et al. *NEJM* (2005) Gluckman P et al. *Lancet* (2005) Azzopardi D et al. *NEJM* (2009)

Simbruner G et al. *Pediatrics* (2010) Jacobs S et al. *Arch Pediatr Adolesc Med* (2011)



### **Potential Therapies to Augment Neuroprotection**

#### Anticonvulsant or antiexcitatory

Phenobarbital, topiramate, levetiracetam, xenon, magnesium sulfate, bumetanide

#### Anti-inflammatory or antioxidant

Sodium cromoglicate, minocycline, indomethacin, melatonin, N-acetylcysteine, allopurinol, pomegranate polyphenols, 7-nitroindazole, 2-iminobiotin, necrostatin 1

#### Multiple mechanisms

Erythropoietin

#### Growth factors and cell-based therapies

Nerve growth factor, insulin-like growth factor 1, brain derived neurotrophic factor, autologous cordblood transplantation

Modified from Johnston MV et al. Lancet Neurology 2011

california perinatal guality care collaborative Modified from Johnston MV et al. Lancet Neurology 2011



# Erythropoietin (Epo)

Generally used for erythropoiesis, may provide neuroprotection

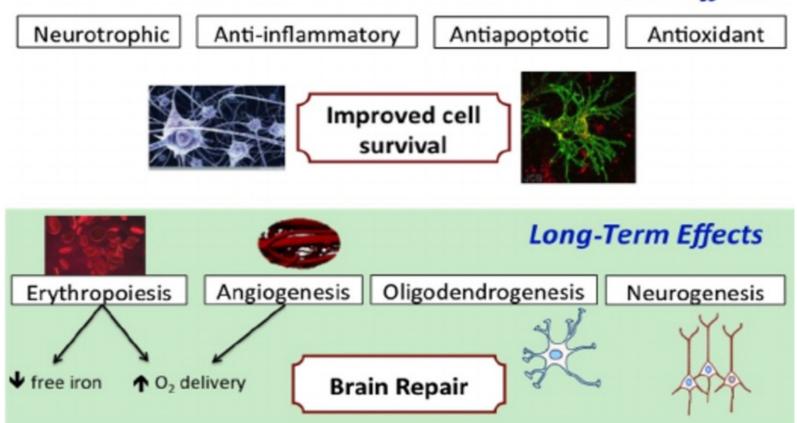


Figure from Juul SE & Pet GC. *Clinics Perinatol* (2015)

california perinatal quality care collaborative

#### **Acute Effects**

### High Dose Epo for Asphyxia and Encephalopathy HEAL trial – Phase III

**Design**: Double masked RCT Epo 1,000 units/kg IV x 5 doses or placebo initiated at <24 hours of age.

**Primary outcome:** Death or moderate to severe neurodevelopmental impairment at 18-24 months of age.

**Secondary outcomes:** Safety, brain injury by MRI at 7 days of age, and serial biomarkers of brain injury.

Sample size: 500 infants, completed enrollment October 2019 and follow-up in October 2021.

PIs: Yvonne Wu, MD MPH and Sandra Juul, MD PhD

### Severity of encephalopathy and outcome

	Mortality	Disability	Normal
	(0/0)	(0/0)	$\left( ^{0\!\! /0}\right)$
Mild	0	0	100
Moderate	5	24	71
Severe	80	20	0
All	13	14	73

Robertson C et al., Dev Med Child Neurol (1985) Thornberg E et al., Acta Pediatrica (1995)



### What is the current practice for MILD encephalopathy?

Therapeutic drift reported in network data and small cohort studies

- Children's Hospital Network –77% of mild HIE patients cooled
- Vermont Oxford Neonatal (VON) Encephalopathy Registry 40% did not meet criteria for moderate-severe NE
- California Perinatal Quality Care Collaborative (CPQCC) 69% of patients with mild HIE cooled

Various definitions of MILD NE are used

- All three registries use the VON definition of NE which is not comparable to trial inclusion criteria
- Some use the presence of 1 abnormality and others use 2 or more of any severity on Sarnat exam

# **Outcome of MILD HIE**

PRIME study –

- Met NICHD criteria and modified Sarnat with  $\geq$  1 abnormality, NOT cooled
- 28 (52%) with abnormal short-term outcome (aEEG n=4, MRI n=9, abnormal discharge exam n=22)
- At 18-22 months disability seen in 7 (16%)
  - 7 with Bayley III Cognitive score <85
  - 1 with cerebral palsy (GMFCS level 4)
  - 2 with autism

Prempunpong C et al. *J Perinatol* 2017 Chalak LF et al. *Ped Res* 2019



# **TIME** trial

#### Lucile Packard Children's Hospital



TIME trial: Therapeutic Hypothermia for Infants with Mild HIE

#### Loma Linda Children's





Rady Children's



CHOC Children's



Benioff Oakland Children's

### CPQCC

### **TIME trial: Specific Aims**

- To determine the best approach to identify, consent, and randomize term newborns with mild HIE.
- To demonstrate that therapeutic hypothermia for mild HIE is safe.
- To demonstrate that therapeutic hypothermia for mild HIE results in improved scores (≥ 1 standard deviation) on the Warner Initial development evaluation of adaptive and functional skills (WIDEA-FS) and the Alberta Infant Motor Scale (AIMS) at 12-14 months of age, as an early marker of neurodevelopmental outcome.



### TIME trial: Study design

- Sample size: 68 infants 34 per treatment arm
- Intervention: 72 hours of TH to 33.5 ± 0.5°C or control 72 hours of targeted temperature Management: 36.5-37.3°C using esophageal or rectal probe
- **Primary Outcome:** WIDEA and AIMS at 12-14 months of age
- Secondary Outcomes: length of stay, need for central lines, intubation, use of sedation, feeding status at discharge, Bayley at 24 months
- Clinical Sites:
  - Lucile Packard Children's Hospital Children's Hospital Orange County (CHOC) Loma Linda Children's Hospital Rady Children's Hospital Benioff Children's Hospital Oakland

### Seizures during rewarming and outcome

**Objective:** Determine incidence of seizures during rewarming and association with abnormal outcome

**Design:** Nested cohort study of infants in the NICHD Optimizing Cooling RCT

**Methods:** aEEG tracings during rewarming were compared to 12-hour period prior to re-warming. Outcomes were determined at 2 years of age

**Results:** Seizures were more common during rewarming (27% versus 14%, p=0.001). Death or disability was more common in those with seizures during rewarming (RR 1.7 95% CI 1.25-2.37)

Chalak LF et al. JAMA Neurol (2021)

### Feeding during cooling

Concerns for NEC limited feedings during cooling in the RCTs, but there may be beneficial effects of enteral feeds after HIE.

A study of National Health Service data (2010-2017) reported 6030 neonates received TH and 31% were fed. NEC seen in 7(0.1%). Using propensity analysis, babies that were fed had less LOS, higher survival to discharge, and shorter NICU stay. Residual confounding could not be ruled out.

A survey of UK NICUs found that the rate of enteral feeding during 2014-2016 was 59%. The number of NICUs feeding increased significantly during this period. There was an increase in NICUs feeding as chronologic age increased. 58% used only donor or expressed BM.

Babies less likely to be fed if had severe HIE, on inotropes, or elevated lactate.

Hazeldine B, et al. *BMJ Paediatr* (2017) Gayle C, et al. *Lancet Child Adol* (2021)

### Anti-seizure medications (ASM) at discharge

The use of AEDs following HIE is controversial.

~10% of neonates with HIE develop epilepsy. Highest risk in those with neonatal seizures and abnormal brain imaging. Of those with neonatal seizures, 25% develop epilepsy. The highest rate epilepsy is with status epilepticus or severe brain injury. Glass H, et al. Ped Res 2011

In a neonatal seizure registry (2015-2018) n=303, 64% had continuation of AEDs after discharge. Both epilepsy rate and functional neurodevelopment were similar. Glass H, et al. JAMA Neurol 2021

**Conclusions:** These results support discontinuation of ASM prior to discharge in most infants with acute seizures.

Glass H et al. *Ped Res* 2011 Glass H et al. *JAMA Neurol* 2021



### **Additional resources**



### Seminars of Fetal and Neonatal Medicine (SFNM) 2021

#### **Issue 1.** Daily management of neonates with NE treated with hypothermia *Guest Editors:* Sonia Bonifacio, Mohamed El-Dib, and Pia Wintermark, on behalf of the Newborn Brain Society

## **Issue 2.** Unanswered questions for neonates with NE treated with hypothermia *Guest Editors:* Sonia Bonifacio, Mohamed El-Dib, and Pia Wintermark, on behalf of the Newborn Brain Society



### **Additional resources**



Weekly webinars on topics related to neonatal neurology organized by the Newborn Brain Society Education Committee

#### April 2021 – Speakers & Topics

- Update on Neonatal Seizures II Hannah Glass, MDCM, MAS and Janet Soul, MDCM, FRCPC
- Neuroimaging I Manon Benders, MD, PhD and Sudhin Thayyil, MD, FRCPCH, PhD
- *Newborn Brain Injury The Global Perspective –* Joy Lawn, BMedSci, MB BS, MRCP(Paeds), MPH, PhD and Cally Tann, MBChB, PhD, FRCPCH
- Parents and Neonatal Brain Care Monica Lemmon, MD and Eleanor Molloy, MB, BCh, BAO, PhD, FRCPCH, FRCPI
- Inter-alpha Inhibitor Proteins, Neuroprotection after Hypoxic Ischemic Brain Injury Barbara Stonestreet, MD

www.newbornbrainsociety.org

### **Additional resources**

### 13TH INTERNATIONAL NEWBORN BRAIN CONFERENCE

...........

FEBRUARY 9-12 • 2022 Clearwater Beach - Florida, United States

Both in-person and virtual

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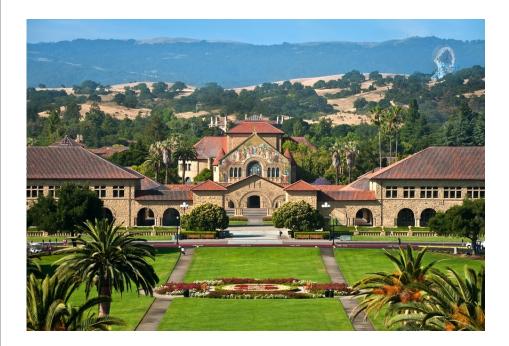


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### Thank you!



### Demo of the CoolTool

Tom Shimotake, MD

## *CoolTool*: https://www.cpqcc.org/cooltool/

### Northern California Neonatal Cooling Consortium Meetings started in 2010

- UCSF Benioff Children's Hospital Mission Bay ICN and Oakland NICU
- Stanford University Children's Hospital NICU
- Santa Clara Valley Medical Center
- Kaiser Permanente Northern CA
- California Pacific Medical Center/Sutter Health System

### Working Group to develop screening toolkit for CPQCC

"Early Screening and Identification of Candidates for Neonatal Therapeutic Hypothermia Toolkit"

- Priya Jegatheesan
- Anna Morgan
- Thomas Shimotake
- Dongli Song
- Krisa Van Meurs

### **Released February 2015**





## *CoolTool*: https://www.cpqcc.org/cooltool/



Mridu Sinha, PhD



University of California, San Diego PhD · Bioengineering and Biomedical Engineering

2012 - 2017

UCSF

#### University of California, San Francisco

Certificate Program · Implementation Science: Translating Evidence into Practice, Policy and Public Health

2016 - 2017

- Developed project idea for decision support tool based on CPQCCC Early Screening Toolkit

- Interviewed staff and providers at multiple birth centers throughout CA
- Multiple iterations of protocol and flowchart based on feedback.
- **Goal:** provide easy to use and openly accessible decision support tool for rapid identification and referral of potential candidates for cooling





Back to Cool Tool Resource Page

#### CoolTool Neuro Analysis

This tool is intended to promote **identification and early referral** of babies at risk for hypoxic-ischemic encephalopathy (HIE). It **does not determine if a baby should be cooled**. The decision to proceed with cooling should only be made after consultation with a Regional Cooling Center CCS numbered letter), based upon their institutional criteria. These recommendations are based on CPQCC's Neonatal Therapeutic Hypothermia toolkit.

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Disclaimer: These suggested guidelines are not a substitute for clinical judgement.

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### **CoolTool** https://www.cpqcc.org/cooltool/

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## *CoolTool* Clinical data field

https://www.cpqcc.org/cooltool/

Enter details below		
Gestational Age >= 35wks *	◯ Yes ◯ No	Inknown
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Age in Hours *		
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Apgar @10min		
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Infant Blood Gas within 1hr	рН	BaseDef
of life		
CPR/Epinephrine	Unknown	~
Continued need for	○ Yes ○ No	unknown
assisted ventilation at 10		
mins of life.		
Change Input	Determine Recon	nmendations >>



### It's 3:13am.....

- You receive a page at 3:13am to attend an emergent c/s delivery
- 38 y.o. G0P1 @ 41w+1d GA
- Severe prolonged decel after SROM w/ cord prolapse
- Mother is prepped in OR and you are ready but await anesthesia...
- Emerg C/S done under GAS. Baby del @ 3:45am (~32min of low HR)
- Baby emerges blue, limp and apneic. HR is <60bpm

### It's 3:13am.....

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- Baby emerges blue, limp and apneic. HR is <60bpm

## Baby emerges blue, limp and apneic...

- You begin resus:  $W/D/S/S \Rightarrow BMV$  i FiO2 to 100%. HR <60bpm
- Despite effective BMV at 100% the **HR <60bpm.** CC begin by 1.5min
- BMV and CC  $\Rightarrow$  intubated by 6min of life.
- Return of HR > 60 bpm so CC held (total 4min of CC). No epi given
- Some color and tone improvement by 8 min of life
- Some resp effort noted and reflex irritability by 12min of life

Apgars	Color	Heart Rate	Resp Effort	Tone	Reflex Irrit	Total
1 min	0	1	0	0	0	1
5 min	0	1	0	0	0	1
10 min	1	2	<b>0</b> (*intub)	1	0	*4
15 min	1	2	<b>1</b> (*intub)	1	1	*6



## Baby emerges blue, limp and apneic...

- You begin resus:  $W/D/S/S \Rightarrow BMV$  FiO2 to 100%. **HR <60bpm**
- CC begin by 1.5min for HR <60bpm despite effective BMV at 100%
- BMV and CC  $\Rightarrow$  intubated by 6min of life.
- Return of HR >60bpm so CC held (total 4min of CC). No epi given
- Some color and tone improvement by 8 min of life
- Some resp effort over vent and reflex irritability noted by 12min of life.

Apgars	Color	Heart Rate	Resp Effort	Tone	Reflex Irrit	Total
1 min	0	1	0	0	0	1
5 min	0	1	0	0	0	1
10 min	1	2	<b>0</b> (*intub)	1	0	*4
15 min	1	2	<b>1</b> (*intub)	1	1	*6



## *CoolTool* Clinical data field

Enter details below		
Gestational Age >= 35wks *	• Yes • No	unknown
2		
Age in Hours *	1	
Acute Perinatal Events	Cord Prolaps	e
	Uterine Rupt	ure
	Severe Fetal	Bradycardia
Apgar @10min	1	
Worst Cord Blood Bas 🛛 👔	рН	BaseDef
Infant Blood Gas within 1hr	рН	BaseDef
of life		
CPR/Epinephrine	CPR	<b>~</b>
Continued need for	• Yes • No	unknown
assisted ventilation at 10		
mins of life.		
Change Input	Determine Recor	nmendations >>

https://www.cpqcc.org/cooltool/

## CoolTool Neonatal Neuro Assessment Tool

Gestational Age >= 35wks \* ( Yes Time of life at neuro exam: 1 Hr No unknown Min ? Seizures: No known episode EEG Confirmed Suspected/Clinical ? Age in Hours \* \*\* Click to select characteristics for each row in the table below Acute Perinatal Events ? Cord Prolapse Lethargic Normal Level of Irritable / Cannot Uterine Rupture Stupor / Obtunded See Conciousness Hyperalert Unresponsive Assess video See video Severe Fetal Bradycardia Apgar @10min ? Normal Jittery / Spontaneous Decreased No Activity Cannot Increased <u>See</u> Activity Assess See video See video Gas 👔 pH Worst Cord Blog <u>video</u> See video BaseDef Infant Blood G within 1hr pН BaseDef Distal Flexion Normal Slight flexion Complete Posture Cannot extension <u>See</u> Decerebrate Extension Assess <u>video</u> See video See video CPR/Epinephrine ? Continued need for Yes No unknown Normal Flaccid Increased Hypotonic Tone Cannot See assisted ventilation at 10 Assess See video See video See video <u>video</u> mins of life. ? Normal Change Input Uncoordinated Weak Absent Suck Cannot See Assess See video See video See video <u>video</u> Normal Incomplete Moro Cannot See Exagerated Absent See video Assess <u>video</u> Cannot Gag Normal Absent Assess Unable to Likely severity: Evaluate Print Assess

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#### CoolTool Neuro Analysis

Enter details below

of life

This tool is intended to promote identification and early referral of babies at risk for hypoxic-ischemic encephalopathy (HIE). It does not determine if a baby should be cooled. The decision to proceed with cooling should only be made after consultation with a Regional Cooling Center CCS numbered letter), based upon their institutional criteria. These recommendations are based on CPQCC's Neonatal Therapeutic Hypothermia toolkit

Neonatal Neuro Tool

Switch to detailed view

Back to Cool Tool Resource Page

## FiO2 weaned, cord gas and baby gas obtained

- HR remains >100bpm with improving color and perfusion
- O2 sats >95% and FiO2 weaned back to 21% by 20min of life
- The baby begins to recovery some tone and spont resp effort
- Cord UA gas sent: 7.25/58/28/-5



## FiO2 weaned, cord gas and baby gas obtained

- HR remains >100bpm with improving color and perfusion
- O2 sats >95% and FiO2 weaned back to 21% by 20min of life
- The baby begins to recovery some tone and spont resp effort
- Cord UA gas sent: 7.25/58/28/-5

\*Does this cord gas mean the baby is not at risk for HIE?

H. Sabir, et.al, J Pediatr, 161 (2012)

## Cord gas and baby gas obtained

- HR remains >100bpm with improving color and perfusion
- O2 sats >95% and FiO2 weaned back to 21% by 20min of life
- The baby begins to recovery some tone and spont resp effort
- Cord UA gas sent: 7.20/52/28/-8
- Patient blood gas obtained at 30min: 6.98/42/-15
- Initial glucose = 28
- IV placed and given 2ml/kg D10W and 10ml/kg NS bolus
- Prepping for UA/UV



## *CoolTool* Neonatal Neuro Assessment Tool

Enter details below	Neonatal Neuro T	ool			Switch to c	detailed viev
Gestational Age >= 35wks * 💿 Yes 🔗 No 🔷 unknown	Time of life at neu	o exam:	🗘 Hr	Min		
2	Seizures: No	known epis	ode EEG (	Confirmed	Suspected/Clinica	al
Age in Hours * 🔋 1	** Click to select of	haracteris	tics for each ro	w in the table l	below	
Acute Perinatal Events 🔐 Cord Prolapse Uterine Rupture Severe Fetal Bradycardia	Level of Conciousness	Normal See video	Irritable / Hyperalert	Lethargic / Obtunded <u>See video</u>	Stupor / Unresponsive	Cannot Assess
Apgar @10min         ?         4           Worst Cord Blood Gas         ?         7.2         -8	Spontaneous Activity	Normal See video	Jittery / Increased See video	Decreased See video	No Activity See video	Cannot Assess
Infant Blood Gas within 1hr 6.98 -15 of life 2 CPR/Epinephrine 2 CPR	Posture	Normal See video	Slight flexion / extension <u>See video</u>	Distal Flexion / Complete Extension <u>See video</u>	Decerebrate	Cannot Assess
Continued need for <u>Yes No unknown</u> assisted ventilation at 10	Tone	Normal See video	Increased See video	Hypotonic See video	Flaccid See video	Cannot Assess
mins of life.	Suck	Normal See video	Uncoordinated See video	Weak See video	Absent See video	Cannot Assess
	Moro	Normal See video	Exagerated	Incomplete See video	Absent	Cannot Assess
	Gag		Normal		sent	Cannot Assess
	Likely reverity:	Unable			Evaluate	Print

This tool is intended to promote identification and early referral of babies at risk for hypoxic-ischemic encephalopathy (HIE). It does not determine if a baby should be cooled. The decision to proceed with cooling should only be made after consultation with a Regional Cooling Center CCS numbered letter),

Back to Cool Tool Resource Page

based upon their institutional criteria. These recommendations are based on CPQCC's Neonatal Therapeutic Hypothermia toolkit

CAOCC

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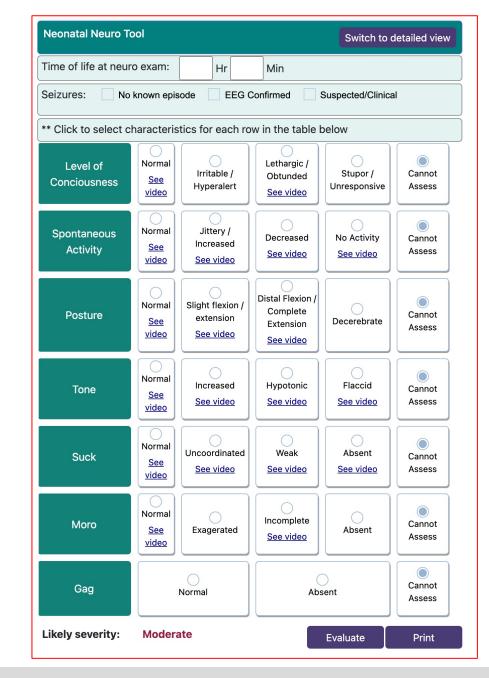
CoolTool Neuro Analysis

https://www.cpqcc.org/cooltool/

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## *CoolTool* Neonatal Neuro Assessment Tool







#### Back to Cool Tool Resource Page

#### CoolTrol Neuro Analysis

This tool is invested to promete the number of the number of babies at risk for hypoxic-ischemic encephalopathy (HIE). It does not determine if a baby should be cooled. The decision to proceed with cooling should only be made after consultation with a Regional Cooling Center CCS numbered letter), based upon their institutional criteria. These recommendations are based on CPQCC's Neonatal Therapeutic Hypothermia toolkit.

## *CoolTool* Neonatal Neuro Assessment Tool

Enter details below			
Gestational Age >= 35wk	S *	🔿 Yes 🔵 No	unknown
	?	[	
Age in Hours *	?		
Acute Perinatal Events	?	Unknown Placental Abr Cord Prolaps	e
Apgar @10min	?		
Worst Cord Blood Gas	?	рН	BaseDef
Infant Blood Gas within 1	nr	рН	BaseDef
of life	?		
CPR/Epinephrine	?	Unknown	~
Continued need for assisted ventilation at 10 mins of life.	?	🔵 Yes 🔵 No	unknown
Change Input	ľ	Determine Recon	nmendations >>

Disclaimer: These suggested guidelines are not a substitute for clinical judgement.

[[]] This work is licensed under a Creative Commons Attribution-NonCommercial-ShareAlike 4.0 International License

https://www.cpqcc.org/cooltool/

## *CoolTool* Neonatal Neuro Assessment Tool

CPQCC Back to Cool Tool Resource Page california perinata quality care collaborative CoolTool Neuro Analysis Neonatal Neuro Tool Time of life at neuro exam: Hr Min No known episode EEG Confirmed Suspected/Clinical Seizures: \*\* Click to select characteristics for each row in the table below Lethargic / Obtunded Normal Level of Conciousness Irritable / Hyperalert Stupor / Unresponsive Cannot Assess See video See video Normal Jittery / Increased Decreased No Activity Spontaneous Activity Cannot Assess See video See video See video See video Slight flexion / Distal Flexion / Normal Posture extension **Complete Extension** Decerebrate Cannot Assess See video See video See video Normal Increased Hypotonic Flaccid Tone Cannot Assess See video See video See video See video Uncoordinated Weak Absent Normal Suck Cannot Assess See video See video See video See video Normal Incomplete Moro Absent Cannot Assess Exagerated See video See video Gag Normal Absent Cannot Assess **Print Analysis** 

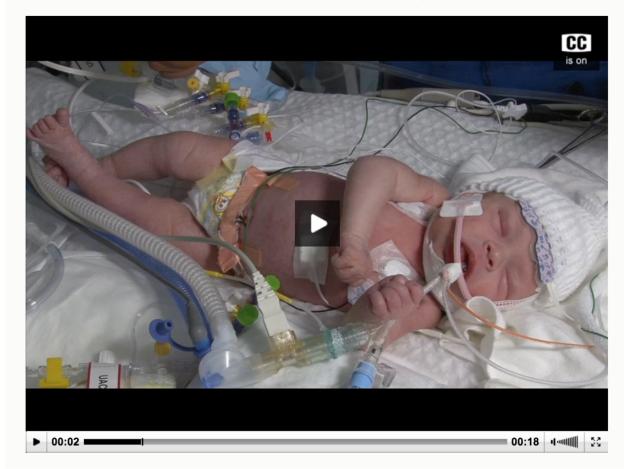
https://www.cpqcc.org/cooltool/

The Neurologic Exam for Neonates with Suspected Encephalopathy

by Courtney Wustoff, MD (Stanford University Lucile Packard Children's Hospital)

### Neurologic Exam: Level of Consciousness Video (Moderate)

Video: Moderate Encephalopathy > Level of consciousness > Lethargic





PediNeurologic Exam: A Neurodevelopmental Approach

by Paul D. Larsen, MD (Univ of Nebraska College Med)

### Suzanne S. Stensaas, PhD (Univ of Nebraska College Med)

#### Site Index

INTRODUCTION
 Developmental Milestones

DEVELOPMENTAL ANATOMY • Major Events & Growth NEWBORN • Normal Exam • Abnormal Exam 3 MONTH OLD • Normal Exam

MONTH OLD
 Normal Exam

12 MONTH OLD • Normal Exam

• Normal Exam

21/2 YEAR OLD • Normal Exam

• For all 6 age groups

SITE CONTENTS

...Listed by Topic

OTHER SECTIONS OF INTEREST • Credits • Copyright

- Contacts & Feedback
- How to Use This Site
- Download Movies Page
- Movie Download Instructions
- Password to Unlock Zipped Movies
- Creative Commons License: Movie Use
- How to Fix Messy Fonts
- How to Show Closed Captions 
  How to add QuickTime to PowerPoint
- How to add Quick Time to PowerPo
- Health Content Disclaimer
   Site Production Notes
- Home



#### PediNeurologic Exam: A Neurodevelopmental Approach

Paul D. Larsen, M.D. The University of Nebraska, College of Medicine





california perinatal quality care collaboratives://neurologicexam.med.utah.edu/pediatric/ht

### Initial Neuro assessment by exam at 1 hour of life

- Baby breathing spontaneously  $\implies$  extubated to CPAP+6@21%.
- Baby is obtunded but moves spontaneously
- Making occasional short, rapid, jittery movements
- Strong distal flexion with closed fists and ankles. Legs extended.
- Hypertonic
- Weak suck, Incomplete Moro, Gag absent
- PERRL, Breathing spontaneously on NCPAP.

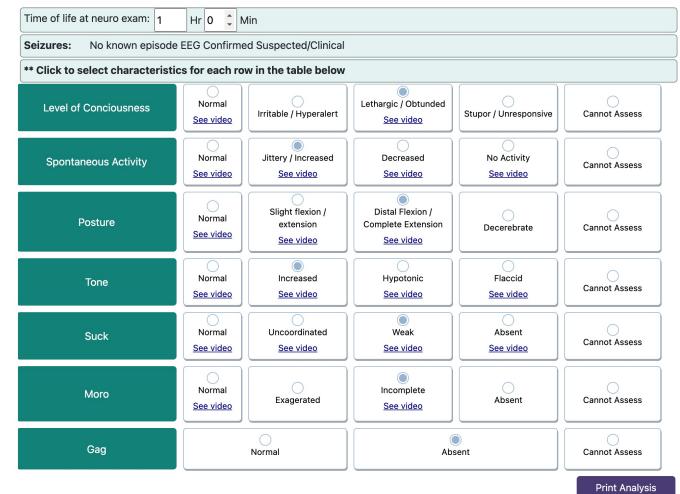


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Back to Cool Tool Resource Page

CoolTool Neuro Analysis

#### **Neonatal Neuro Tool**



## *CoolTool*: Neonatal Neuro Assessment Tool

https://www.cpqcc.org/cooltool/



Back to Cool Tool Resource Page

#### CoolTool Neuro Analysis

This tool is intended to promote **identification and early referral** of babies at risk for hypoxic-ischemic encephalopathy (HIE). **It does not determine if a baby should be cooled**. The decision to proceed with cooling should only be made after consultation with a Regional Cooling Center CCS numbered letter), based upon their institutional criteria. These recommendations are based on CPQCC's Neonatal Therapeutic Hypothermia toolkit.

## *CoolTool* Neonatal Neuro Assessment Tool

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#### Enter details below

Gestational Age >= 35wks *	Yes No unknown
?	
Age in Hours *	1
Acute Perinatal Events 🛛 👔	Cord Prolapse Uterine Rupture Severe Fetal Bradycardia
Apgar @10min	4
Worst Cord Blood Gas 🛛 👔	7.2 8
Infant Blood Gas within 1hr	6.98
of life	
CPR/Epinephrine	CPR 🗸
Continued need for	🔵 Yes 🔵 No 🔵 unknown
assisted ventilation at 10	
mins of life.	
Change Input	Determine Recommendations >>

#### Recommendations

#### AT RISK.

#### Suggested Actions:

Refer the table below for suggested action based on neurologic assessment. For Cooling, consider the **worst exam** after initial resuscitation

Neurologic Assessment Result	Clinical Suggestions
Normal	Screens negative at <i>this</i> time. Symptoms may change. You may perform subsequent neuro exam using <u>NeuroTool</u> . <b>Continue to monitor</b> as per <u>CPQCC guidelines</u> .
Mild	Call cooling center to discuss case. Provide care as per the management guidelines for potential candidates
Moderate to Severe	Call cooling center to discuss the need for transfer and cooling. Provide care as per the management guidelines <u>for potential</u> <u>candidates</u>
Incomplete Exam	Call cooling center to discuss case immediately.

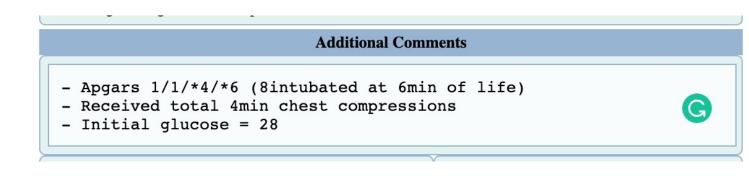
#### Find Regional Cooling Center

See protocols related to cooling: Choose a protocol: Go Print Report



## CoolTool Printable report with - time stamp (age)

- Additional comments
- MD signature Date and time



	Paitent History			
Gestational Age	>= 35 Weeks			
Age in hours	1			
Sentinal events	Cord Prolapse; Severe Fetal Bradycardia			
Apgar @10 mins	4			
Worst cord gas	7.2; 8			
Infant cord gas	6.98; 15			
CPR/Epinephrine	CPR			
Continued need for assisted vent @10 mins	Yes			
	Recommendations			
AT RISK.				
Perform neurologic assessment after resu	scitation.			
	Neuro Evaluation			
Time of Life at Neuro Exam	1 Hr 0 Min			
Seizures	No known Episode			
Level of Consciousness	Lethargic/Obtunded			
Spontaneous Activity	Decreased			
Posture	Distal Flexion/Complete Extension			
Tone	Normal/Increased			
Suck	Cannot Assess			
Moro	Cannot Assess			
Gag	Cannot Assess			
Like	ly Neuro Assessment Result			
Moderate to Severe: Call cooling center t the management guidelines for potential	to discuss the need for transfer and cooling.Provide care as per candidates			
	Additional Comments			
<ul> <li>Apgars 1/1/*4/*6 (8intul</li> <li>Received total 4min ches</li> <li>Initial glucose = 28</li> </ul>				
MD Signature	Date/Time			

https://www.cpqcc.org/cooltool/

## Other resource documents on

## CoolTool.info





Back to Cool Tool Resource Page

#### CoolTool Neuro Analysis

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#### Enter details below

Gestational Age >= 35wks *	● Yes ○ No ○ unknown
?	
Age in Hours *	1
Acute Perinatal Events 👔	Cord Prolapse Uterine Rupture Severe Fetal Bradycardia
Apgar @10min	4
Worst Cord Blood Gas 🛛 👔	7.2 8
Infant Blood Gas within 1hr	6.98 15
of life	
CPR/Epinephrine	CPR 🗸
Continued need for	🔵 Yes 🔵 No 🔵 unknown
assisted ventilation at 10	
mins of life.	
Change Input	Determine Recommendations >>

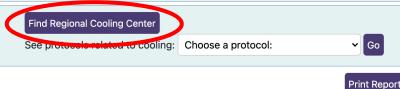
#### Recommendations

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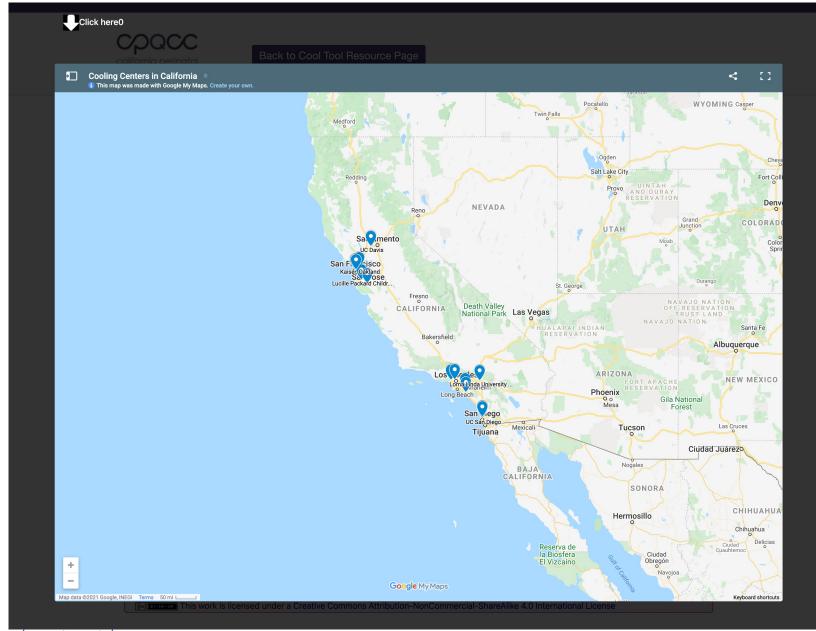
CoolTool

Regional Cooling

Center finder

https://www.cpqcc.org/cooltool/

### CPQCC



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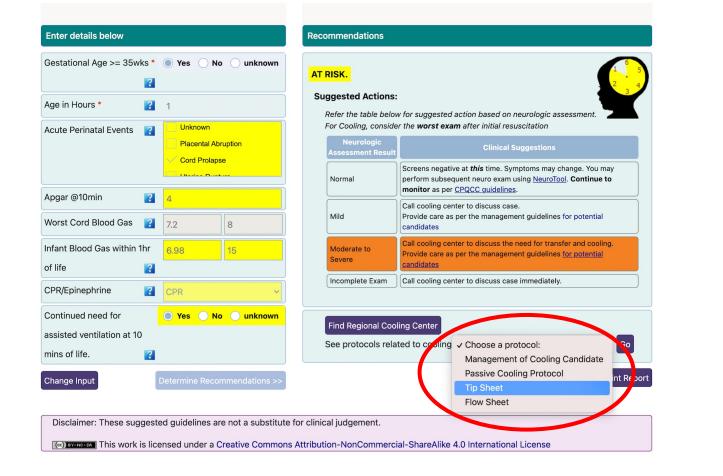
X





#### CoolTool Neuro Analysis

This tool is intended to promote **identification and early referral** of babies at risk for hypoxic-ischemic encephalopathy (HIE). **It does not determine if a baby should be cooled**. The decision to proceed with cooling should only be made after consultation with a Regional Cooling Center CCS numbered letter), based upon their institutional criteria. These recommendations are based on CPQCC's Neonatal Therapeutic Hypothermia toolkit.



## *CoolTool* Protocols and flow sheets

https://www.cpqcc.org/cooltool/

## Protocols and flow sheets

Vital Signs Record														
DOB: Time: : 1 <sup>ST</sup> Temperature after Birth: Date: Time:									Time:	Appendix D				
Radiant Warmer turned off :  No  Yes: Date: Time::									;	<b>Guidelines for Passive Cooling</b>				
Ordered By Doctor:MD														
	Baseline	15"	30"	45"	1 hr 1h	1. Document Regional Cooling Center contacted and decision made to initiate passive coo								
Date		to be a candidate for cooling.												
Time	me ABBREVIATED TIP SHEET FOR COOLTOOL													
Axillary	°C	-									Appendix E			
HR RR	1) Complete and document Norma Assessment (cooltecting)											info)	<b>Lanagement of Screened Neonates Who Do Not Qualify for Cooling</b> I neonates who meet screening criteria will require or qualify for cooling therapy. However,	
BP													hay still have significant risks factors that warrant special consideration. These risks may from mild acidosis to multi-organ dysfunction. In addition, initial signs of neonatal halopathy may be subtle and neurologic symptoms may evolve over time. In some cases,	
	aintain ba ature: Eve	11	3) Turn down/off external heat sources						urces				e cooling may already have been initiated. Patients without clinical evidence of perinatal injury should be rewarmed only after a thorough evaluation and consultation (phone/video) neonatologist at a regional cooling center. Levels of concern and need for observation or interventions/therapies may be appropriate depending upon the clinical presentation.	
	e: on adm		) Monitor core (rectal) temperature											
	a. Target rectal temperature = 33.5°C (92.									° <b>C</b> (92.	2.3°F) Maintain communication with regional cooling center			
b. Check temp 15 min and record on flow								and re	cord or	n flow s	low sheet.		<ul> <li>a. Discuss management and plan if significant clinical changes develop.</li> <li>If heat sources were removed/cooling was initiated, slowly begin rewarming</li> </ul>	
https://www.cpqcc.org/cooltool/													<ul> <li>a. Document time of lowest temperature and source (e.g., axillary vs. rectal).</li> <li>b. Rewarm with target rate of approximately 0.5 °C /hour. Avoid overheating.</li> </ul>	
										3			<ul> <li>Monitor temperature periodically</li> <li>a. Target rectal/core temp = 36.5°C (97.7°F) or axillary/skin temp = 36.0°C</li> </ul>	
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#### **Key Collaborators**

- Mridu Singha (UCSD)
- Sonia Bonifacio (Stanford/LPCH)
- Courtney Wutsoff (Stanford/LPCH)
- Krisa Van Meurs (Stanford/LPCH)
- Anna Morgan (Kaiser, Oakland)
- Joseph Schulman (CA DHHS/CA Children's Services
- Dongli Song (SCVMC)
- Priscilla Joe (UCSF BCH Oakland)
- Alexis Davis (Stanford/LPCH)
- Alex Espinoza (Alta Bates Medical Center)

### **Bay Area Cooling Coolborative**

- UCSF BCH Oakland Children's Hospital
- Stanford Unversity/Lucille Packard Children's Hospital
- Northern California Kaiser/Oakland
- Santa Clara Valley Medical Center
- CPMC/Sutter San Francisco Medical Center
- UC Davis Medical Center

### **CPQCC PQUIP**

- Henry Lee (Stanford/LPCH)
- Priya Jeegathesan (SCVMC)

### **UCSF NICN TEAM**

- Donna Ferriero
- Hannah Glass
- Fernando Gonzalez
- Yao Sun
- Mark Petersen
- Dawn Gano
- Jim Barkovich
- NICN Nursing Staff

### Our Patients and Families





Demo of NeoCool

Mary Harbert, MD

### **Question and Answer Session**

Moderated by: Henry Lee, MD & Priya Jeegatheesan, MD

### **SPEAKERS:**

Dongli Song, MD, PhD Krisa Van Meurs, MD Tom Shimotake, MD Mary Harbert, MD



Closing

# Big thanks to our speakers & moderators and thank YOU for attending this webinar.



